

Claims

1. A kit comprising:

a first article having a surface;

a peptide sequence immobilized relative to or adapted to be immobilized relative

to the surface, the peptide sequence including a portion of a cell surface receptor that
interacts with an activating ligand such as a growth factor to promote cell proliferation,
the portion including enough of the cell surface receptor to interact with the activating
ligand and the portion free of interchain binding region to the extent necessary to prevent
spontaneous binding between portions; and

a candidate drug for affecting the ability of the peptide sequence to bind to other
identical peptide sequences in the presence of the activating ligand.

2. A kit as in claim 1, further comprising a second article having a surface and the
peptide sequence immobilized relative to or adapted to be immobilized relative to the
surface of the second article.

3. A kit as in claim 1, wherein the peptide sequence is MGFR.

4. A method comprising:

providing a peptide including a portion of a cell surface receptor that interacts
with an activating ligand such as a growth factor to promote cell proliferation, the
portion including enough of the cell surface receptor to interact with the activating ligand
and the portion free of interchain binding region to the extent necessary to prevent
spontaneous binding between portions;

exposing the peptide to a candidate drug for affecting the ability of the activating
ligand to interact with the peptide, and to the activating ligand; and

determining the ability of the candidate drug to prevent interaction of the
activating ligand with the peptide.

5. A method as in claim 4, comprising determining the ability of the candidate drug
to prevent interaction of the peptide with other proteins or peptides.

6. A method as in claim 4, comprising providing a first article having a surface and a plurality of the peptides immobilized relative to or adapted to be immobilized relative to the surface;

exposing the peptides and the surface of the first article to the candidate drug and
5 at least one activating ligand; and
determining the ability of the candidate drug to prevent interaction of the activating ligand with the peptide.

7. A method as in claim 4, comprising providing a first article having a surface, a
10 second article having a surface, and a plurality of the peptides immobilized relative to or adapted to be immobilized relative to the surfaces of the first and second articles;

exposing the peptides and the surfaces of the first and second articles to the candidate drug and at least one activating ligand; and
15 determining immobilization of the first and second articles relative to each other.

8. A method as in claim 4, wherein the step of exposing the peptides to the candidate drug and at least one activating ligand comprises exposing the peptide and the candidate drug to one or both of cell lysate and cell supernatant containing the activating
20 ligand.

9. A method as in claim 4, wherein the peptide sequence is PSMGFR.

10. A method of treating a subject to reduce the risk of or progression of cancer comprising:

25 administering to a subject who is known to be at risk for cancer or is diagnosed with cancer an agent for inhibiting interaction of an activating ligand with a portion of a cell surface receptor that interacts with the activating ligand to promote cell proliferation.

11. A method as in claim 10, comprising administering to the subject an agent for
30 inhibiting inductive multimerization of the portion of the cell surface receptor that interacts with the activating ligand to promote cell proliferation.

12. A method as in claim 10, wherein the cell surface receptor is MUC1

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13. A method as in claim 10, wherein the portion of the cell surface receptor is MGFR.

5 14. A method as in claim 10, wherein the portion of the cell surface receptor contains a significant part of the PSMGFR sequence.

15. A method as in claim 10, comprising administering to the subject an agent for inhibiting dimerization of the portion of the cell surface receptor that interacts with the
10 activating ligand to promote cell proliferation.

16. The method of claim 10, wherein the cancers is selected from the group consisting of: breast, prostate, lung ovarian, colorectal, and brain cancer.

15 17. A method as in claim 10, wherein the activating ligand is a multimer.

18. The method of claim 10, wherein the activating ligand is a protein with a molecular weight of about 17kD.

20 19. The method of claim 10, wherein the activating ligand is a protein with a molecular weight of about 23kD.

20. The method of claim 10, wherein the activating ligand is a protein with a molecular weight of about 35kD.

25 21. The method of claim 10, wherein the activating ligand contains sequences derived from the protein 14-3-3.

22. The method of claim 10, wherein the activating ligand contains sequences
30 derived from cathepsin D.

23. The method of claim 10, wherein the activating ligand contains sequences derived from NM23.

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24. The method of claim 10, wherein the activating ligand contains sequences derived from human annexin V.

5 25. The method of claim 10, wherein the activating ligand contains sequences derived from beta-lipotropin.

26. The method of claim 10, wherein the activating ligand is a cleavage product of proopiomelanocortin.

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27. The method of claim 10, wherein the portion of the cell surface comprises at least 12 contiguous amino acids from the sequence
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVSDVPPFSAQSGA.

15 28. The method of claim 10, wherein the portion of the cell surface receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region comprises at least 12 contiguous amino acids from the peptide sequence
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVS.

20 29. The method of claim 10, wherein the agent is selected for use in the method by determining its ability to bind to a significant portion of the peptide,
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVSDVPPFSAQSGA.

30. The method of claim 10, wherein the agent is selected for use in the method by
25 determining its ability to bind to a significant portion of the peptide sequence
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVS.

31. A method of treating a subject to reduce the risk or of progression of cancer comprising:

30 administering to a subject who is known to be at risk of cancer or is diagnosed with cancer, an agent for preventative clustering of portions of cell surface receptors that interact with an activating ligand such as a growth factor to promote cell proliferation.

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32. The method of claim 31, wherein the cell surface receptor is MUC1.

33. A method as in claim 31, comprising contacting the portions of the cell surface
5 receptor with a molecule that can bind to multiple portions thereby clustering a plurality
of the portions.

34. A method as in claim 31, wherein the portion is MGFR

10 35. A method as in claim 31, wherein the portion contains a significant amount of the
PFMGFR sequence.

36. The method of claim 31, wherein the cancer is selected from the group consisting
of: breast, prostate, lung ovarian, colorectal, and brain cancer.

15 37. The method of claim 31, wherein the portion of the cell surface receptor
comprises at least 12 contiguous amino acids from the peptide sequence
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVSDVPFPFSAQSGA.

20 38. The method of claim 31, wherein the portion of the cell surface receptor
comprises at least 12 contiguous amino acids from the peptide sequence
GTINVHDTVETQFNQYKTEAASPYNLTISDVSVS.

39. The method of claim 31, wherein the specific binding portion of the agent is
25 selected for use in the method by determining its ability to bind to a significant portion of
the peptide, GTINVHDTVETQFNQYKTEAASPYNLTISDVSVSDVPFPFSAQSGA.

40. The method of claim 31, wherein the specific binding portion of the agent is
selected for use in the method by determining its ability to bind to a significant portion of
30 the peptide, GTINVHDTVETQFNQYKTEAASPYNLTISDVSVS.

41. A kit comprising:

a species able to become immobilized relative to a shed cell surface receptor interchain binding region; and

a signaling entity immobilized relative to or adapted to be immobilized relative to the species.

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42. A kit as in claim 41, wherein the species binds to a portion of a shed MUC1 receptor that is connected to the interchain binding region.

43. A kit as in claim 41, wherein the cell surface receptor is MUC1.

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44. The kit as in claim 41, wherein the signaling entity is a colloid particle.

45. The kit as in claim 41, wherein the signaling entity is not a colloid particle.

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46. The colloid particle as in claim 41, further comprising a colloid particle, wherein the signaling entity is attached to the colloid particle.

47. A composition comprising:

at least a portion of a shed cell surface receptor interchain binding region; and

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a signaling entity immobilized relative to or adapted to be immobilized relative to the portion.

48. A kit comprising:

a species able to bind to a portion of a cell surface receptor that remains attached

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to the cell surface after shedding of a cell surface receptor interchain binding region; and

a signaling entity immobilized relative to or adapted to be immobilized relative to the species.

49. A kit as in claim 48, wherein the cell surface receptor is MUC1.

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50. The kit as in claim 48, wherein the signaling entity is a colloid particle.

51. The kit as in claim 48, wherein the signaling entity is not a colloid particle.

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52. The kit particle as in claim 48, further comprising a colloid particle, wherein the signaling entity is attached to the colloid particle.

5 53. The kit as in claim 48, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 17kD.

10 54. The kit as in claim 49, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 23kD.

15 55. The kit as in claim 49, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 35kD

20 56. The kit as in claim 49, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains sequences derived from the protein 14-3-3

57. The kit as in claim 49, wherein the portion comprises 14-3-3.

58. The kit as in claim 49, wherein the portion comprises cathepsin D.

25 59. The kit as in claim 49, wherein the portion comprises NM23.

60. The kit as in claim 49, wherein the portion comprises Human annexin V.

30 61. The kit as in claim 49, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains at least one sequence derived from beta-lipotropin.

62. The kit as in claim 49, wherein the species able to bind to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a cleavage product of proopiomelanocortin..

5 63. A kit comprising:

a species able to bind to a portion of a cell surface receptor that includes the interchain binding region; and

a signaling entity immobilized relative to or adapted to be immobilized relative to the species.

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64. A kit as in claim 63, wherein the cell surface receptor is MUC1.

65. The kit as in claim 63, wherein the signaling entity is a colloid particle.

15 66. The kit as in claim 63, wherein the signaling entity is not a colloid particle.

67. The colloid particle as in claim 63, further comprising a colloid particle, wherein the signaling entity is attached to the colloid particle.

20 68. A peptide species comprising:

at least a fragment of a sequence that corresponds to that portion of a cell surface receptor that interacts with an activating ligand such as a growth factor to promote cell proliferation, the portion being detached from any cell; and

an affinity tag.

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69. A peptide species as in claim 68, wherein the affinity tag is connected to the fragment.

30 70. A peptide species as in claim 68, wherein the affinity tag defines a portion of a continuous amino acid sequence that includes both the fragment and the affinity tag.

71. The species of claim 68, wherein the affinity tag is a polyamino acid tag.

72. The species of claim 68, wherein the affinity tag is a polyhistidine tag.

73. The species of claim 68, wherein the affinity tag is a GST tag.

5 74. The species of claim 68, wherein the affinity tag is biotin.

75. The species of claim 68, wherein the affinity tag is Thioredoxin.

76. The species of claim 68, wherein the affinity tag is selected to bind to a species
10 immobilized with respect to the surface of an article.

77. The species of claim 68, further comprising an article having a surface, and a
species able to capture the affinity tag immobilized with respect to the surface.

15 78. The species of claim 68, wherein the article is a particle.

79. The species of claim 68, wherein the affinity tag is fastened to the C-terminus of
the portion of the receptor.

20 80. The species of claim 68, wherein the cell surface receptor is MUC1.

81. The species of claim 68, wherein the cell surface receptor portion comprises 12 or
more contiguous amino acids in the sequence

GTINVHDTVETQFNQYKTEAASPYNLTISDVSVSDVPFPSAQSGA (SEQ ID NO: 7)

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82. A peptide species as in claim 68, wherein the fragment comprises at least a
portion of PSMGFR.

83. A peptide species as in claim 68, wherein the fragment comprises PSMGFR.

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84. A peptide species as in claim 68, wherein the fragment comprises at least a
fragment of the sequence that corresponds to that portion of MUC1 that interacts with an

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activating ligand such as a growth factor to promote cell proliferation in association with MUC1-dependent tumorigenesis.

85. A peptide species as in claim 68, wherein the fragment comprises enough of the sequence that corresponds to that portion of MUC1 that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region in association with MUC1-dependent tumorigenesis such that a biomolecule that interacts with that portion of MUC1 that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region in association with MUC1-dependent tumorigenesis interacts with the fragment.

86. A kit comprising:
a particle; and
at least a fragment of the sequence that corresponds to that portion of a cell surface receptor that interacts with an activating ligand such as a growth factor to promote cell proliferation, the fragment being detached from any cell, fastened to or adapted to be fastened to the particle.

87. The kit of claim 86, wherein the cell surface receptor is MUC1.

88. A kit comprising:
an article having a surface; and
a biomolecule that binds to a portion of a cell surface receptor that interacts with an activating ligand such as a growth factor to promote cell proliferation, the biomolecule being fastened to or adapted to be fastened to the surface of the article.

89. The kit of claim 88, wherein the article comprises a particle.

90. The kit of claim 88, wherein the cell surface receptor is MUC1.

91. The kit of claim 88, further comprising:
a second particle; and

a portion of a cell surface receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region, the portion being detached from any cell, fastened to or adapted to be fastened to the second particle.

5 92. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 17kD.

10 93. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 23kD.

15 94. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a protein of about 35kD.

20 95. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains sequences derived from the protein 14-3-3.

96. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains sequences derived from Cathepsin D.

25 97. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains sequences derived from NM23.

30 98. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains sequences derived from human annexin V.

99. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region contains at least one sequence derived from beta-lipotropin.

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100. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is a cleavage product of proopiomelanocortin.

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101. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region is selected from the group which includes calcimycin, fusaric acid, L- α -methyl-dopa, and etomoxir.

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102. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region comprises calcimycin.

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103. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region comprises fusaric acid.

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104. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region comprises L- α -methyl-dopa.

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105. The kit as in claim 88, wherein the biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region comprises etomoxir.

106. The composition of claim 88, wherein the biomolecule is derived from a cell line selected from the group consisting of HTB-133, CRL-1504, and CRL-1500.

107. A method comprising:

exposing a ligand capable of binding with a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region, and an agent capable of blocking said binding, to a candidate drug for
5 disruption of interaction between the ligand and the agent; and
determining disruption of the interaction by the candidate drug.

108. A method comprising:

10 exposing a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region which is capable of binding with a ligand, and an agent capable of blocking said binding, to a candidate drug for disruption of interaction between the portion and the agent; and
determining disruption of the interaction by the candidate drug.

15 109. A method comprising:

exposing a synthetic drug, and a biological target of the synthetic drug, to a candidate drug which may interact with the biological target to a degree greater than the interaction between the synthetic drug and the target; and
determining disruption of the interaction by the candidate drug.

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110. A method as in claim 109, wherein the synthetic drug is a derivative of fusaric acid.

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111. A method as in claim 109, wherein the synthetic drug is a derivative of L- α -methyl-dopa.

112. A method as in claim 109, wherein the synthetic drug is a derivative of etomoxir.

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113. A method for treating a subject having a cancer characterized by the aberrant expression of MUC1, comprising:
administering to the subject fusaric acid in an amount effective to reduce tumor growth.

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121. A method as in claim 119, wherein the method comprises administering to the subject etomoxir in an amount effective to block the interaction of a natural ligand and the portion of the MUC1 receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region.

122. A method as in claim 119, wherein the method comprises administering to the subject etomoxir in an amount effective to reduce shedding of the interchain binding region of the MUC1 receptor.

5 123. The method of 119, wherein the levels of shed interchain binding region are reduced relative to a level measured in a past sample.

124. The method of 119, wherein the levels of shed interchain binding region are reduced relative to a control sample.

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125. A method for treating a subject having a cancer characterized by the aberrant expression of MUC1, comprising:
administering to the subject L- α -methyl-dopa in an amount effective to reduce tumor growth.

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126. A method as in claim 125, wherein the subject is otherwise free of symptoms calling for treatment with L- α -methyl-dopa.

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127. A method as in claim 125, wherein the method comprises administering to the subject L- α -methyl-dopa in an amount effective to block the interaction of a natural ligand and the portion of the MUC1 receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region.

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128. A method as in claim 125, wherein the method comprises administering to the subject L- α -methyl-dopa in an amount effective to reduce shedding of the interchain binding region of the MUC1 receptor.

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129. The method of 125, wherein the levels of shed interchain binding region are reduced relative to a level measured in a past sample.

130. The method of 125, wherein the levels of shed interchain binding region are reduced relative to a control sample.

131. A method for treating a subject having a cancer characterized by the aberrant expression of MUC1, comprising:
administering to the subject calcimycin in an amount effective to reduce tumor growth.

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132. A method as in claim 131, wherein the subject is otherwise free of symptoms calling for treatment with calcimycin.

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133. A method as in claim 131, wherein the method comprises administering to the subject calcimycin in an amount effective to block the interaction of a natural ligand and the portion of the MUC1 receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region.

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134. A method as in claim 131, wherein the method comprises administering to the subject calcimycin in an amount effective to reduce shedding of the interchain binding region of the MUC1 receptor.

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135. The method of 134, wherein the levels of shed interchain binding region are reduced relative to a level measured in a past sample.

136. The method of 134, wherein the levels of shed interchain binding region are reduced relative to a control sample.

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137. A method for treating a subject having a cancer characterized by the aberrant expression of MUC1, comprising:
administering to the subject butylindazole in an amount effective to reduce tumor growth.

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138. A method as in claim 137, wherein the subject is otherwise free of symptoms calling for treatment with butylindazole.

139. A method as in claim 137, wherein the method comprises administering to the subject butylindazole in an amount effective to block the interaction of a natural ligand

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and the portion of the MUC1 receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region.

140. A method as in claim 137, wherein the method comprises administering to the
5 subject butylindazole in an amount effective to reduce shedding of the interchain binding region of the MUC1 receptor.

141. The method of 137, wherein the levels of shed interchain binding region are reduced relative to a level measured in a past sample.

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142. The method of 137, wherein the levels of shed interchain binding region are reduced relative to a control sample.

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143. A method for treating a subject having a cancer characterized by the aberrant expression of MUC1, comprising:

administering to the subject NS1619 in an amount effective to reduce tumor growth.

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144. A method as in claim 143, wherein the subject is otherwise free of symptoms calling for treatment with NS1619.

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145. A method as in claim 143, wherein the method comprises administering to the subject NS1619 in an amount effective to block the interaction of a natural ligand and the portion of the MUC1 receptor that remains attached to the cell surface after shedding of the cell surface receptor interchain binding region.

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146. A method as in claim 143, wherein the method comprises administering to the subject NS1619 in an amount effective to reduce shedding of the interchain binding region of the MUC1 receptor.

147. The method of 143, wherein the levels of shed interchain binding region are reduced relative to a level measured in a past sample.

148. The method of 143, wherein the levels of shed interchain binding region are reduced relative to a control sample.

149. A method comprising:

5 exposing a composition selected among calcimycin, butylindazole, NS1619, fusaric acid, L- α -methyl-dopa, and etomoxir, and a biomolecule that binds to a portion of a cell surface receptor that remains attached to the cell surface after shedding of a cell surface receptor interchain binding region, to a candidate drug which may interfere with interaction between the composition and the biomolecule; and

10 determining disruption of the interaction by the candidate drug.

150. A method of treating a subject having cancer or at risk for developing cancer comprising:

15 administering to the subject an agent that reduces cleavage of a cell surface receptor.

151. A method of treating a subject having cancer or at risk for developing cancer comprising:

20 administering to the subject an agent that reduces cleavage of a cell surface receptor interchain binding region from the cell surface.

152. The method of claim 150, wherein the cell surface receptor is MUC1.

25 153. The method of claim 151, wherein the interchain binding region comprises a contiguous amino acid sequence of at least 12 amino acids from the sequence GFLGLSNIKFRPGSVVVQLTLAFRE.

30 154. The method of claim 150, wherein the interchain binding region comprises a contiguous amino acid sequence of about 12 to 18 amino acids, within the region of the human MUC1 receptor amino acids 507 through 549 (refers to Spicer et al sequence – corresponds to amino acids 1067 through 1100 of Genbank accession # PI5941, PID G547937).

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evaluating indication of cancer or potential for cancer based upon the determining step.

163. The method of claim 158, wherein the interchain binding region comprises a
5 contiguous amino acid sequence of at least 12 amino acids from the sequence
GFLGLSNIKFRPGSVVVQLTLAFRE.

164. The method of claim 158, wherein the interchain binding region comprises a
contiguous amino acid sequence of about 12 to 18 amino acids, within the region of the
10 human MUC1 receptor amino acids 507 through 549 (refers to Spicer et al sequence –
corresponds to amino acids 1067 through 1100 of Genbank accession # PI5941, PID
G547937).

165. The method of claim 158, wherein the interchain binding region comprises a
15 contiguous amino acid sequence of about 12 to 18 amino acids, within the region of the
human MUC1 receptor amino acids 525 through 549 (refers to Spicer et al sequence –
corresponds to amino acids 1085 through 1109 of Genbank accession # PI5941, PID
G547937).

20 166. The method of claim 160, wherein the sample is a fluid sample.

167. The method of claim 160, wherein the sample is blood.

168. The method of claim 160, wherein the sample is a tissue sample.

25 169. The method of claim 160, wherein the sample is a proliferating cell line derived
from a subject's cells.

170. The method of claim 158, wherein the cancer is characterized by aberrant
30 expression of MUC1.

171. The method of claim 158, wherein the amount of interchain binding region is
determined by a method selected from the group consisting of MALDI, western blotting,

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PCR, LCR, rtPCR, cycling probe technology, gel electrophoresis, or antibody-based assay, magnetic cell sorting, fluorescence activated cell sorting, bead-based assays or an ELISA assay.

- 5 172. The method of claim 158, wherein the amount of interchain binding region is determined by an aggregation assay.

173. The method of claim 158, wherein the amount of interchain binding region is determined by a colloid-based method such as colloid-colloid or colloid-bead assay.

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174. The method of claim 158, wherein the sample is selected from the group consisting of: a needle biopsy, a tissue specimen, a tissue surface in an intraoperative procedure, and a tissue surface or cellular solution in a minimally invasive procedure such as a laparoscopy.

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175. A method comprising:
 determining a site of cleavage of a cell surface receptor in a sample from a subject; and
 evaluating an indication of cancer or potential for cancer based upon the
20 determining step.

176. The method of claim 175, wherein the cell surface receptor is MUC1.

25 177. The method of claim 175, wherein the sample is selected from the group consisting of: a needle biopsy, a tissue specimen, a tissue surface in an intraoperative procedure, and a tissue surface or cellular solution in a minimally invasive procedure such as a laparoscopy.

178. The method of claim 175, wherein the sample is a fluid sample.

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179. The method of claim 175, wherein the sample is blood.

180. The method of claim 175, wherein the sample is a tissue sample.

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181. The method of claim 175, wherein the cancer is selected from the group consisting of: breast, prostate, lung, ovarian, colorectal, and brain cancer.

5 182. A method as in claim 175, wherein the cancer is characterized by the aberrant expression of MUC1.

183. The method of claim 175, wherein the site of cleavage is determined by a method selected from the group consisting of MALDI, western blotting, PCR, LCR, 10 rtPCR, cycling probe technology, gel electrophoresis, or antibody-based assay, magnetic cell sorting, fluorescence activated cell sorting, bead-based assays or an ELISA assay.

184. The method of claim 175, wherein the amount of interchain binding region is determined by a colloid-based method such as colloid-colloid or colloid-bead assay. 15

185. A method of determining a cleavage site of a cell surface comprising:
contacting a cell with an agent that binds specifically to one potential cell surface
receptor cleavage site and another agent that binds specifically to another potential cell
surface receptor cleavage site; and
20 comparing the ratio of binding of the two agents to the cell surface.

186. The method of claim 185, wherein the surface cell receptor is MUC1.

187. A method of diagnosing a physiological state indicative of cancer or potential for 25 cancer, comprising determining a specific cleavage state of MUC 1 distinguishable from a different cleavage state of MUC1.

188. A method comprising:
determining a first amount of cleavage of a cell surface receptor interchain
30 binding region from a cell surface of a sample from a subject;
determining a second amount of cleavage of a cell surface receptor interchain
binding region from a cell surface of a sample from the subject;
comparing the first amount to the second amount.

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189. A method as in claim 188, comprising comparing the first amount to the second amount as an indication of progression of and/or effectiveness of treatment for cancer.

5 190. A method as in claim 188, comprising comparing the first amount to the second amount as an indication for administration of an agent for prevention of cancer.

191. A method as in claim 188, wherein the subject is undergoing treatment for cancer, the method comprising

10 comparing the first amount to the second amount as an indication of effectiveness of the treatment.

192. A method as in claim 188, wherein the cell surface receptor is MUC1.

15 193. The method of claim 192, wherein the interchain binding region comprises a contiguous amino acid sequence of at least 12 amino acids from the sequence GFLGLSNIKFRPGSVVVQLTLAFRE.

20 194. The method of claim 192, wherein the interchain binding region comprises a contiguous amino acid sequence of about 12 to 18 amino acids, within the region of the human MUC1 receptor amino acids 507 through 549 (refers to Spicer et al sequence – corresponds to amino acids 1067 through 1100 of Genbank accession # PI5941, PID G547937).

25 195. The method of claim 188, wherein the interchain binding region comprises a contiguous amino acid sequence of about 12 to 18 amino acids, within the region of the human MUC1 receptor amino acids 525 through 549 (refers to Spicer et al sequence – corresponds to amino acids 1085 through 1109 of Genbank accession # PI5941, PID G547937).

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196. The method of claim 188, wherein the sample is a fluid sample.

197. The method of claim 188, wherein the sample is blood.

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198. The method of claim 188, wherein the sample is a tissue sample.

199. The method of claim 188, wherein the cancer is selected from the group
5 consisting of: breast, prostate, lung, ovarian, colorectal, and brain cancer.

200. The method of claim 188, wherein the sample is a proliferating cell line derived
from a patient's cells.

10 201. The method of claim 188, wherein the amount of interchain binding region is
determined by a method selected from the group consisting of MALDI, western blotting,
PCR, LCR, rtPCR, cycling probe technology, gel electrophoresis, or antibody-based
assay, magnetic cell sorting, fluorescence activated cell sorting, bead-based assays or an
ELISA assay.

15 202. The method of claim 188, wherein the amount of interchain binding region is
determined by an aggregation assay.

20 203. The method of claim 188, wherein the amount of interchain binding region is
determined by a colloid-based method such as colloid-colloid or colloid-bead assay.

204. The method of claim 188, wherein the sample is selected from the group
consisting of: a needle biopsy, a tissue specimen, a tissue surface in an intraoperative
procedure, and a tissue surface or cellular solution in a minimally invasive procedure
25 such as a laparoscopy.

205. The method of claim 188, wherein the amount of interchain binding region is
determined by a colloid-based method such as colloid-colloid or colloid-bead assay.

30 206. The method of claim 188, wherein the sample is selected from the group
consisting of: a needle biopsy, a tissue specimen, a tissue surface in an intraoperative
procedure, and a tissue surface or cellular solution in a minimally invasive procedure
such as a laparoscopy.

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207. A method as in claim 188, comprising determining a first amount of a cell surface
receptor interchain binding region at the surface of a cell in a sample from a subject,
determining a second amount of a cell surface receptor interchain binding region
5 at the surface of a cell in a sample from the subject,
comparing the first amount to the second amount.

208. A method as in claim 188, comprising determining a first amount of a shed cell
surface receptor interchain binding region in a sample from a subject,
10 determining a second amount of a shed cell surface receptor interchain binding
region in a sample from the subject,
comparing the first amount to the second amount.

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